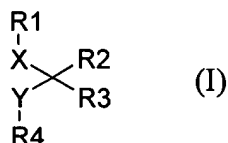


In the claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

1. (currently amended) A compound of Formula I



or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt,

wherein:

~~R<sub>1</sub> is selected from the group consisting of:~~

~~— C<sub>1</sub>-C<sub>6</sub> alkyl, substituted with one or more basic groups;~~

~~— cycloalkyl, substituted with one or more basic groups;~~

~~— aromatic heterocyclyl, comprising at least one nitrogen~~

~~atom, and substituted with one or more basic groups; aliphatic~~

~~heteterocyclyl, comprising at least one nitrogen atom;~~

~~heterocyclyl, comprising at least one hetero atom selected~~

~~from S or O, and substituted with one or more basic groups;~~

~~and~~

~~— aryl, pyridyl substituted with one or more basic groups;~~

R<sub>2</sub> is selected from the group consisting of H, acyl, acylamino,

alkyl, alkylcarbamoyl, alkylthio, alkoxy, aroyl,

aroylamino, aryloxy, arylthio, amidino, amino, aryl,

carbamoyl, carboxy, cyano, cycloalkyl, formyl, guanidino,

halogen, heterocyclyl, hydroxy, oxo, nitro, thiol, a

Z<sub>2</sub>N-CO-O- group, a ZO-CO-NZ- group, and a Z<sub>2</sub>N-CO-NZ- group;

R<sub>3</sub> is selected from the group consisting of COOR<sub>5</sub>, SO(OR<sub>5</sub>), SO<sub>3</sub>R<sub>5</sub>,

P=O(OR<sub>5</sub>)<sub>2</sub>, B(OR<sub>5</sub>)<sub>2</sub>, P=OR<sub>5</sub>(OR<sub>5</sub>), tetrazole, and a carboxylic

acid isostere which is an acidic group having a pKa of from about -5 to about 25;

R<sub>4</sub> is SH, S-CO-C<sub>1</sub>-C<sub>6</sub> alkyl, or S-CO-aryl;

R<sub>5</sub> is H, C<sub>1</sub>-C<sub>6</sub> alkyl, or aryl;

R<sub>6</sub> is H or C<sub>1</sub>-C<sub>6</sub> alkyl;

X is selected from the group consisting of O, S, SO, SO<sub>2</sub>, C(Z)<sub>2</sub>, N(Z), NR<sub>6</sub>SO<sub>2</sub>, SO<sub>2</sub>NR<sub>6</sub>, NR<sub>6</sub>CO, and CONR<sub>6</sub>;

Y is C(Z)<sub>2</sub>; and

Z is independently selected from the group consisting of H, C<sub>1</sub>-C<sub>6</sub> alkyl, aryl, cycloalkyl, and heterocyclyl.

2. (currently amended) The compound according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, wherein:

~~R<sub>1</sub> is selected from the group consisting of:~~

~~— cycloalkyl, substituted with one or more basic groups;~~

~~— heterocyclyl, comprising at least one nitrogen atom;~~

~~heterocyclyl, comprising at least one hetero atom selected from S or O, and substituted with one or more basic groups;~~  
~~and~~

~~— aryl, substituted with one or more basic groups;~~

R<sub>2</sub> is selected from the group consisting of H, acyl, acylamino, alkyl, alkylcarbamoyl, alkylthio, alkoxy, aroyl, aroylamino, aryloxy, arylthio, amidino, amino, aryl, carbamoyl, carboxy, cyano, cycloalkyl, formyl, guanidino, halogen, heterocyclyl, hydroxy, oxo, nitro, thiol, Z<sub>2</sub>N-CO-O-, ZO-CO-NZ-, and Z<sub>2</sub>N-CO-NZ-;

R<sub>3</sub> is COOR<sub>5</sub>;

R<sub>4</sub> is SH, S-CO-C<sub>1</sub>-C<sub>6</sub> alkyl, or S-CO-aryl;

R<sub>5</sub> is H, C<sub>1</sub>-C<sub>6</sub> alkyl, or aryl;

R<sub>6</sub> is H or C<sub>1</sub>-C<sub>6</sub> alkyl;

X is selected from the group consisting of O, S, SO, SO<sub>2</sub>, C(Z)<sub>2</sub>, N(Z), NR<sub>6</sub>SO<sub>2</sub>, SO<sub>2</sub>NR<sub>6</sub>, and CONR<sub>6</sub>;

Y is C(Z)<sub>2</sub>; and

Z is independently selected from the group consisting of H, C<sub>1</sub>-C<sub>6</sub> alkyl, aryl, cycloalkyl and heterocyclyl.

3. (currently amended) The compound according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, wherein:

~~R<sub>1</sub> is selected from the group consisting of:~~

~~—cycloalkyl, substituted with one or more basic groups;~~

~~—heterocyclyl, comprising at least one nitrogen atom; and~~

~~—heterocyclyl, comprising at least one hetero atom selected from S or O, and substituted with one or more basic groups;~~

R<sub>2</sub> is selected from the group consisting of H, C<sub>1</sub>-C<sub>3</sub> alkyl, amino, halogen, and hydroxy;

R<sub>3</sub> is COOR<sub>5</sub>;

R<sub>4</sub> is SH, S-CO-C<sub>1</sub>-C<sub>6</sub> alkyl, or S-CO-aryl;

R<sub>5</sub> is H, C<sub>1</sub>-C<sub>6</sub> alkyl, or aryl;

X is C(Z)<sub>2</sub>;

Y is C(Z)<sub>2</sub>; and

Z is independently H or C<sub>1</sub>-C<sub>6</sub> alkyl.

4. (currently amended) The compound according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, wherein:

~~R<sub>1</sub> is selected from the group consisting of:~~

~~—cycloalkyl, substituted with one or more basic groups; and~~

~~—heterocyclyl, comprising at least one nitrogen atom;~~

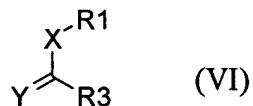
$R_2$  is H, F, or  $C_1$  alkyl;  
 $R_3$  is  $COOR_5$ ;  
 $R_4$  is SH, S-CO- $C_1$ - $C_6$  alkyl, or S-CO-aryl;  
 $R_5$  is H,  $C_1$ - $C_6$  alkyl, or aryl;  
 $X$  is  $C(Z)_2$ ;  
 $Y$  is  $C(Z)_2$ ; and  
 $Z$  is independently H or  $C_1$ - $C_6$  alkyl.

5. (currently amended) The compound according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, wherein:

~~$R_4$  is selected from the group consisting of cyclopentyl, pyridyl, pyrimidinyl, piperidinyl, and thiazolyl;~~

$R_2$  is H, F, or  $C_1$  alkyl;  
 $R_3$  is  $COOR_5$ ;  
 $R_4$  is SH;  
 $R_5$  is H;  
 $X$  is CHZ;  
 $Y$  is CHZ; and  
 $Z$  is independently H or  $C_1$ - $C_6$  alkyl.

6. (Previously presented) A process for the preparation of a compound according to claim 1, wherein  $X$  is  $C(Z)_2$ , and  $R_2$  is H, comprising the step of:  
 reacting a compound of Formula VI,

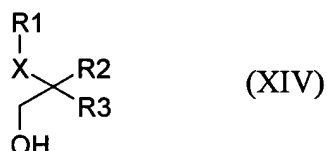


wherein  $R_1$ ,  $R_3$  and  $Y$  are as defined in claim 1 and  $X$  is  $C(Z)_2$ ,  
 with a compound of Formula IX,



wherein  $R_5$  is a protecting group, optionally in the presence of a base or a free-radical initiator.

7. (Previously presented) A process for the preparation of a compound according to claim 1, wherein Y is  $CH_2$ , and X is O, S,  $C(Z)_2$ , or  $N(Z)$ , comprising the step of:  
reacting a compound of Formula XIV,

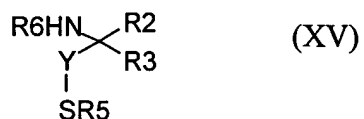


wherein  $R_1$ ,  $R_2$ , and  $R_3$  are as defined in claim 1, and X is O, S,  $C(Z)_2$ , or  $N(Z)$ , with a compound of general Formula IX,



wherein  $R_5$  is a protecting group, in the presence of a suitable reagent, under standard conditions.

8. (Previously presented) A process for the preparation of a compound according to claim 1, wherein X is  $NR_6CO$  or  $NR_6SO_2$ , comprising the step of:  
reacting a compound of Formula XV,



wherein  $R_2$ ,  $R_3$ ,  $R_6$  and Y are as defined in claim 1 and  $R_5$  is a protecting group, with a compound of Formula XVI,

R1-X (XVI)

wherein R<sub>1</sub> is as defined in claim 1 and X is COOH or SO<sub>2</sub>Cl, in the presence of a coupling reagent, under standard conditions.

9. (Previously presented) A pharmaceutical formulation comprising a compound according to any one of claims 1 to 5 as active ingredient in combination with a pharmaceutically acceptable adjuvant, diluent or carrier.

10. (cancelled)

11. (cancelled)

12. (currently amended) A method for treatment or prophylaxis of ~~conditions associated with~~ a condition in which inhibition of carboxypeptidase U is required or desired, comprising administering to a patient in need of such treatment an effective amount of a compound according to any one of claims 1-5.

13. (currently amended) A pharmaceutical formulation for the treatment or prophylaxis of ~~conditions associated with~~ a condition in which inhibition of carboxypeptidase U is required or desired, comprising a compound according to any one of claims 1-5 in combination with a pharmaceutically acceptable adjuvant, diluent, or carrier.

14. (currently amended) A pharmaceutical formulation, comprising:

(i) a compound of Formula I according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt; and

(ii) one or more antithrombotic agents ~~with a different mechanism of action from that of component (i)~~ selected from the group consisting of an antiplatelet agent, thromboxane receptor inhibitor, synthetase inhibitor, fibrinogen receptor antagonist, prostacyclin mimetic, phosphodiesterase inhibitor, and an ADP-receptor (P<sub>2</sub>T) antagonist,

in admixture with a pharmaceutically acceptable adjuvant, diluent, or carrier.

15. (currently amended) A kit of parts comprising:

(i) a pharmaceutical formulation comprising a compound of Formula I according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, in admixture with a pharmaceutically acceptable adjuvant, diluent, or carrier; and

(ii) a pharmaceutical formulation comprising one or more antithrombotic agents ~~with a different mechanism of action from that of component (i)~~ selected from the group consisting of an antiplatelet agent, thromboxane receptor inhibitor, synthetase inhibitor, fibrinogen receptor antagonist, prostacyclin mimetic, phosphodiesterase inhibitor, and an ADP-receptor (P<sub>2</sub>T) antagonist,

in admixture with a pharmaceutically acceptable adjuvant, diluent, or carrier,

wherein compound (i) and agent (ii) are each formulated for administration in conjunction with the other.

16. (currently amended) A method for treatment of a patient suffering from, or susceptible to, a condition in which both

inhibition of carboxypeptidase U and a different antithrombotic mechanism are required or desired, which method comprises administering to the patient a therapeutically effective total amount of:

- (i) a compound of Formula I according to claim 1, or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt, in admixture with a pharmaceutically acceptable adjuvant, diluent, or carrier; and
- (ii) one or more antithrombotic agents ~~with a different mechanism of action from that of component (i)~~ selected from the group consisting of an antiplatelet agent, thromboxane receptor inhibitor, synthetase inhibitor, fibrinogen receptor antagonist, prostacyclin mimetic, phosphodiesterase inhibitor, and an ADP-receptor (P<sub>2</sub>T) antagonist, in admixture with a pharmaceutically acceptable adjuvant, diluent, or carrier.

17. (currently amended) A method for the treatment of a patient suffering from, or susceptible to, a condition in which both inhibition of carboxypeptidase U and a different antithrombotic mechanism are required or desired, which method comprises administering to the patient the formulation according to claim 14.

18. (currently amended) The compound according to any one of claims ~~1-4~~ 1-5, wherein the basic group is selected from the group consisting of amino, amidino, and guanidino.

19. (Previously presented) The process according to claim 6, wherein the protecting group is selected from the group



consisting of acetate (Ac), benzoyl (Bz), benzyl (Bn), and 4-methoxybenzyl (PMB).

20. (Previously presented) The process according to claim 6, wherein the base is selected from the group consisting of NaOMe, NaH, and triethylamine.
21. (Previously presented) The process according to claim 6, wherein the free-radical initiator is  $\alpha,\alpha'$ -azoisobutyronitrile (AIBN).
22. (Previously presented) The process according to claim 7, wherein the protecting group is acetate (Ac) or benzoyl (Bz).
23. (Previously presented) The process according to claim 7, wherein the reagent is  $\text{PPh}_3$ /diisopropyl azodicarboxylate (DIAD).
24. (Previously presented) The process according to claim 8, wherein the protecting group is selected from the group consisting of acetate (Ac), benzoyl (Bz), benzyl (Bn), and 4-methoxybenzyl (PMB).
25. (Previously presented) The process according to claim 8, wherein the coupling reagent is selected from the group consisting of:
  - (i) (benzotriazol-1-yloxy)tripyrrolidinophosphonium hexafluorophosphate (PyBOP)/diisopropylethylamine (DIPEA);
  - (ii) dicyclohexylcarbodiimide (DCC)/1-hydroxybenzotriazol (HOBt);
  - (iii) 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide (EDC)/triethylamine (TEA)/N,N-dimethyl amino pyridine

(DMAP); and

(iv) pyridine.

26-28. (cancelled)